

R^{12} is hydrogen or a group that with $O-R^{11}$ group and with C/4" carbon atom forms a $>C=O$ or epoxy group;

(vi) R^2 is hydrogen, hydroxy, OR^p or alkoxy ;

(vii) A is hydrogen or methyl;

(viii) B is methyl or epoxy;

(ix) E is hydrogen or halogen;

R^3 is hydroxy, OR^p , alkoxy or R^3 is a group that with R^5 and with C/11 and C/12 carbon atoms forms a cyclic carbonate or carbamate, or if W or Z is $>N-R_N$ R^3 is a group that with W or Z forms a cyclic carbamate;

(xi) R^4 is C_1-C_4 alkyl;

(xii) R^5 is hydrogen, hydroxy, OR^p , C_1-C_4 alkoxy, or a group that with R^3 and with C/11 and C/12 carbon atoms forms a cyclic carbonate or carbamate;

(xiii) R^6 is hydrogen or C_1-C_4 -alkyl; and

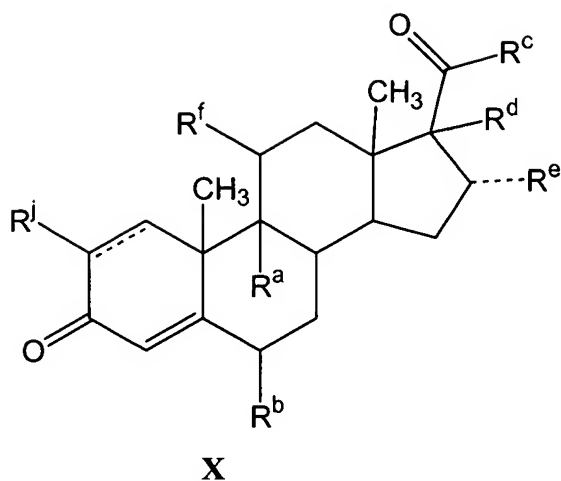
R^p is hydroxyl or amino protective group;

wherein M has a linkage site through which it is linked to S *via* linking group L; provided that the linkage site being is at one or more of the following:

- any reactive hydroxy, nitrogen, or epoxy group located on S^1 , S^2 , or an aglycone oxygen if S^1 and/or S^2 is cleaved off;
- a reactive $>N-R_N$ or $-NR_tR_s$ or oxo group located on Z or W;
- a reactive hydroxy group located at any one of R^1 , R^2 , R^3 , and R^5 ;
- any other group that can be first derivatized to a hydroxy or $-NR_tR_s$ group and

~~R^p is hydroxyl or amino protective group~~

S represents a group of Formula X:



wherein

R^a and R^b independently represents, hydrogen or halogen;

R^c is hydroxy, alkoxy, alkyl, thiocarbonyl, carbonyl or a valence-bond;

R^d and R^e independently represents: hydrogen, hydroxy, methyl or C₁-C₄-alkoxy or each are a group that forms a 1,3-dioxolane ring with the other or a valence bond;

R^f is hydrogen, hydroxy, chloro, or forming a keto group with the carbon atom it is attached to;

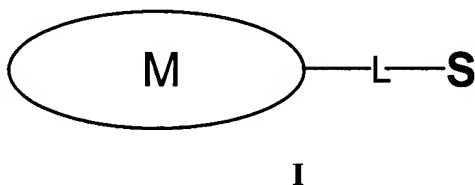
R^j is hydrogen or halogen;

or a pharmaceutically acceptable salt or solvate thereof;

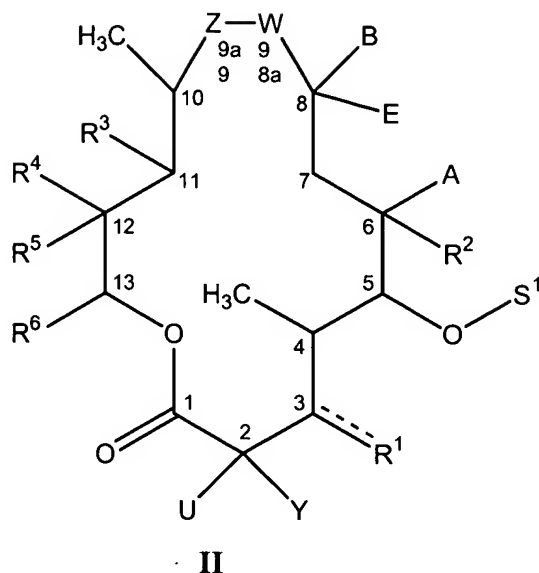
wherein

L is a linker molecule to which each of M and S are covalently linked.

2. (Currently Amended) A compound of the Formula I:



wherein **M** represents a group of Formula II:



wherein

(i) Z and W independently are $>C=O$, $>CH_2$, $>CH-NR_tR_s$, $>N-R_N$ or $>C=N-R_M$, wherein

R_t and R_s independently are hydrogen or alkyl;

R_M is hydroxy, alkoxy, substituted alkoxy or OR^p ;

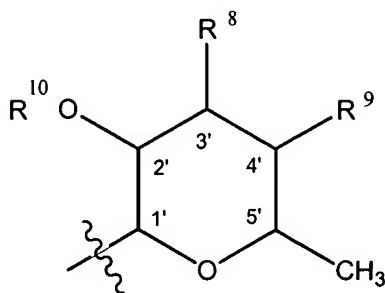
R_N is hydrogen, R^p , alkyl, alkenyl, alkynyl, alkoxy, alkoxyalkyl, or $-C(X)-NR_tR_s$; wherein X is $=O$ or $=S$;

provided that Z and W cannot both simultaneously be, $>C=O$, $>CH_2$, $>CH-NR_tR_s$, $>N-R_N$, $>C=N-R_M$ or a bond;

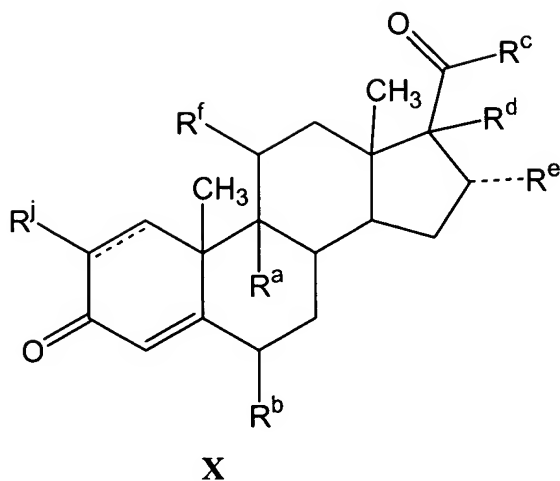
(ii) U and Y independently are hydrogen, halogen, alkyl, or hydroxyalkyl;

(iii) R^1 is hydroxy, OR^p , $-OS^2$ group or an $=O$;

(iv) S^1 is a sugar moiety of Formula **III**:



III



wherein

R^a and R^b independently represents, hydrogen or halogen;

R^c is hydroxy, alkoxy, alkyl, thiocarbamoyl, carbamoyl or a valence-bond;

R^d and R^e independently represents: hydrogen, hydroxy, methyl or C_1 - C_4 -alkoxy or each are a group that forms a 1,3-dioxolane ring with the other or a valence bond;

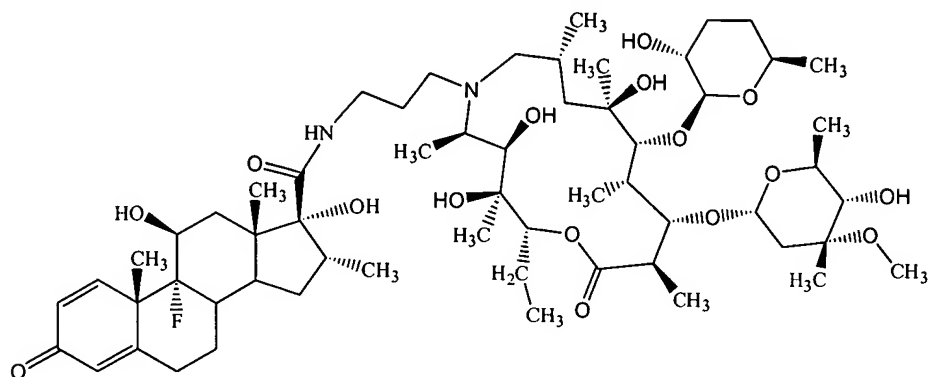
R^f is hydrogen, hydroxy, chloro, or forming a keto group with the carbon atom it is attached to;

R^j is hydrogen or halogen;

or a pharmaceutically acceptable salt or solvate thereof.

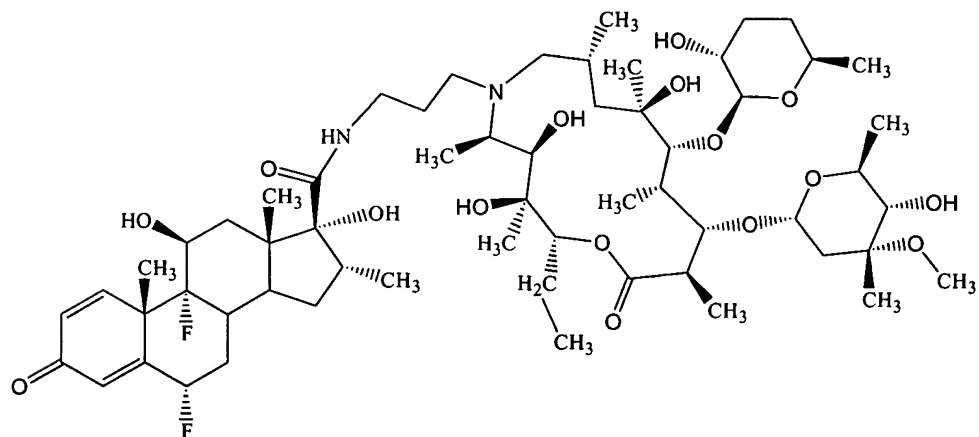
3. (Canceled).
4. (Canceled)
5. (Previously presented) The compound according to claim 2 wherein
 - Z is $>NR_N$, wherein R_N is hydrogen or a methyl group;
 - W is $>CH_2$;
 - B is methyl;
 - E is hydrogen;
 - R^2 is hydroxy;
 - A is methyl;

8. (Previously Presented) A compound of the formula



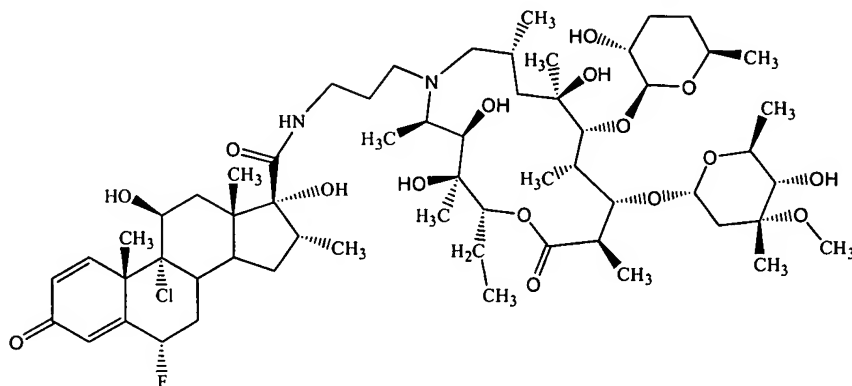
or a pharmaceutically acceptable salt or solvate thereof.

9. (Previously Presented) A compound of the formula



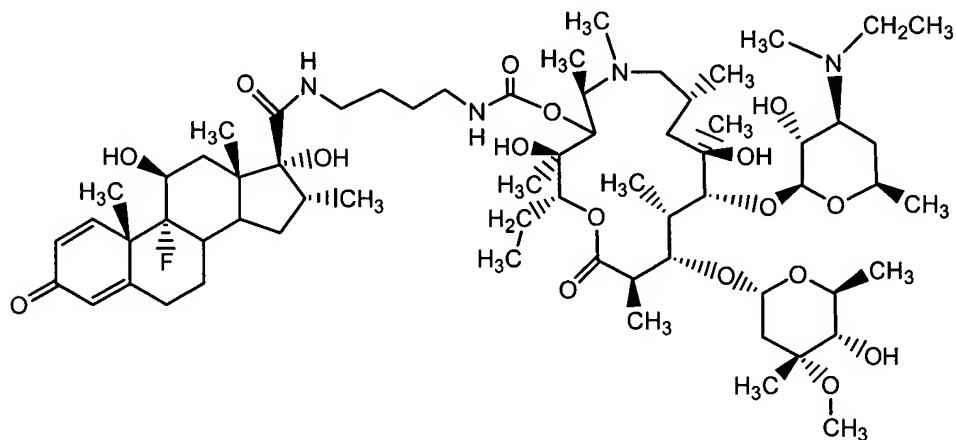
or a pharmaceutically acceptable salt or solvate thereof.

10. (Previously Presented) A compound of the formula



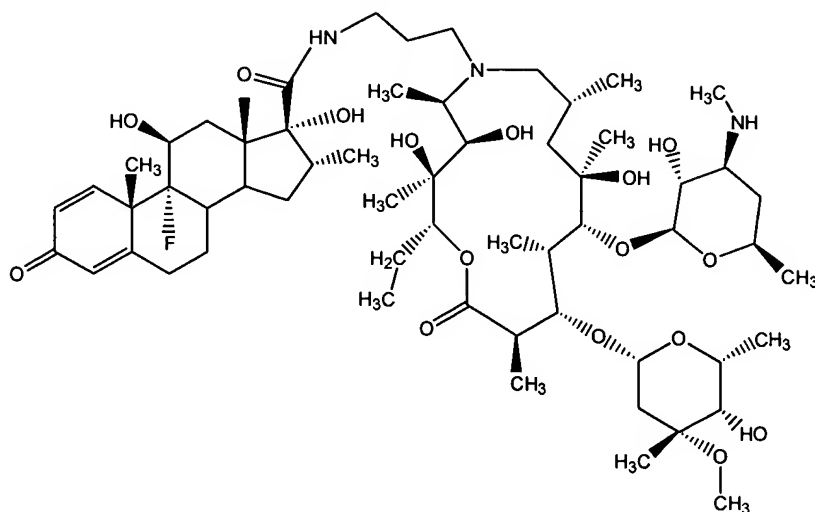
or a pharmaceutically acceptable salt or solvate thereof.

11. (Previously Presented) A compound of the formula



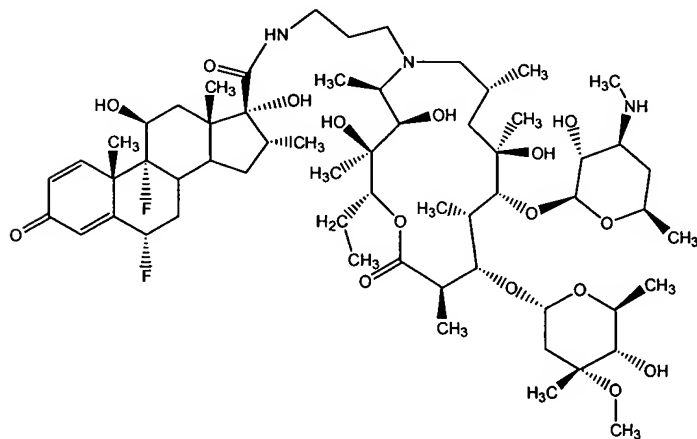
or a pharmaceutically acceptable salt or solvate thereof.

12. (Previously Presented) A compound of the formula



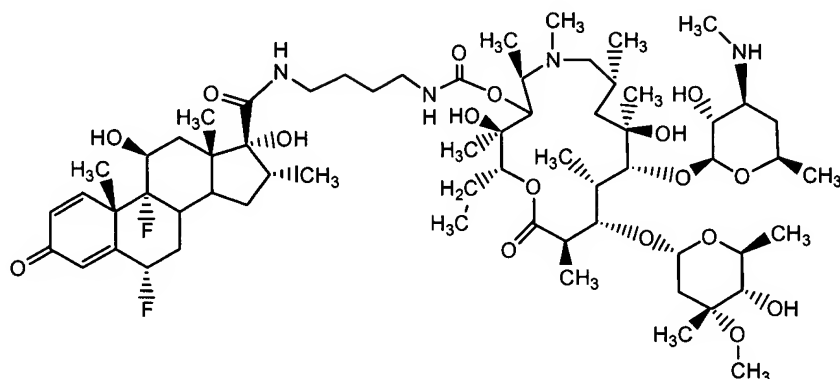
or a pharmaceutically acceptable salt or solvate thereof.

13. (Previously Presented) A compound of the formula



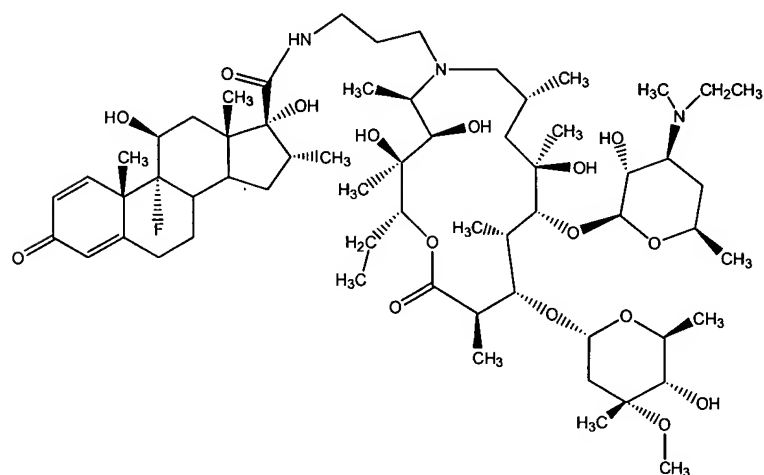
or a pharmaceutically acceptable salt or solvate thereof.

14. (Previously Presented) A compound of the formula



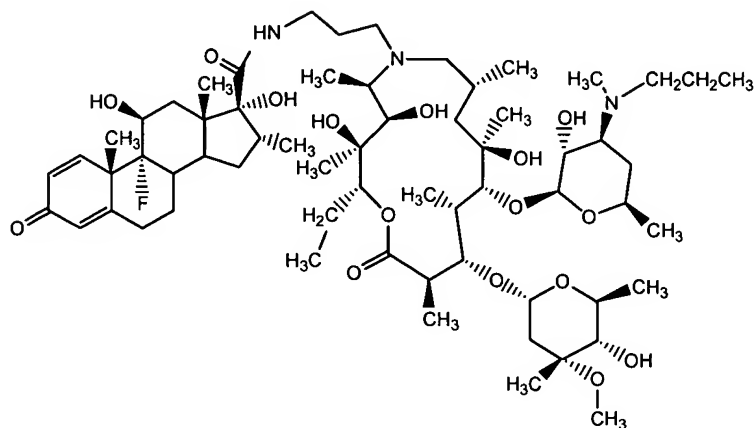
or a pharmaceutically acceptable salt or solvate thereof.

15. (Previously Presented) A compound of the formula



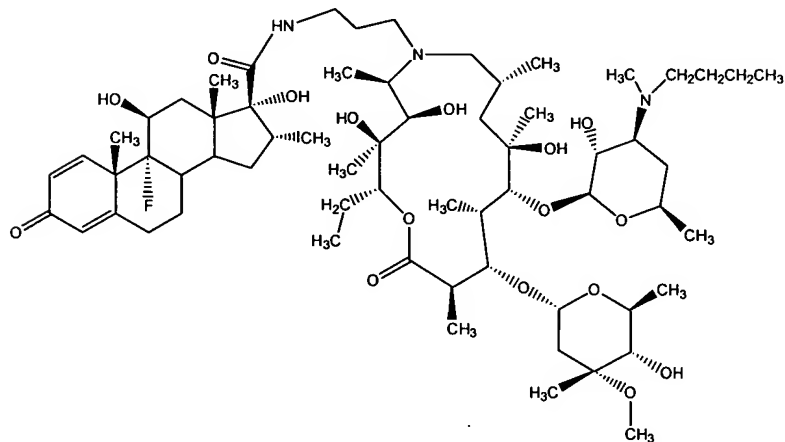
or a pharmaceutically acceptable salt or solvate thereof.

16. (Previously Presented) A compound of the formula



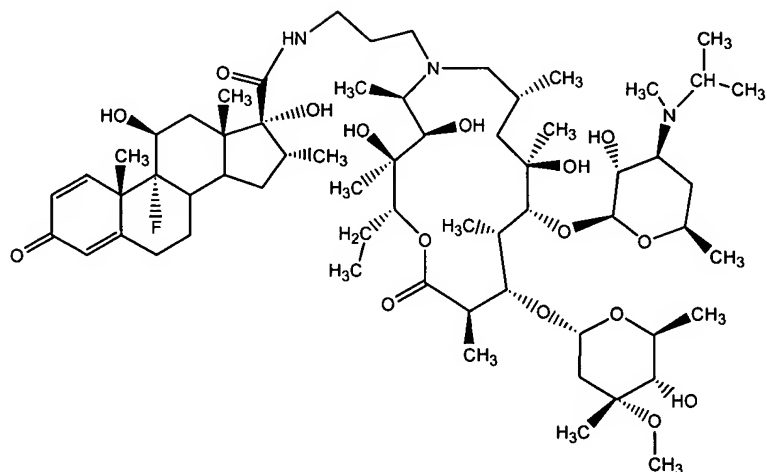
or a pharmaceutically acceptable salt or solvate thereof..

17. (Previously Presented) A compound of the formula



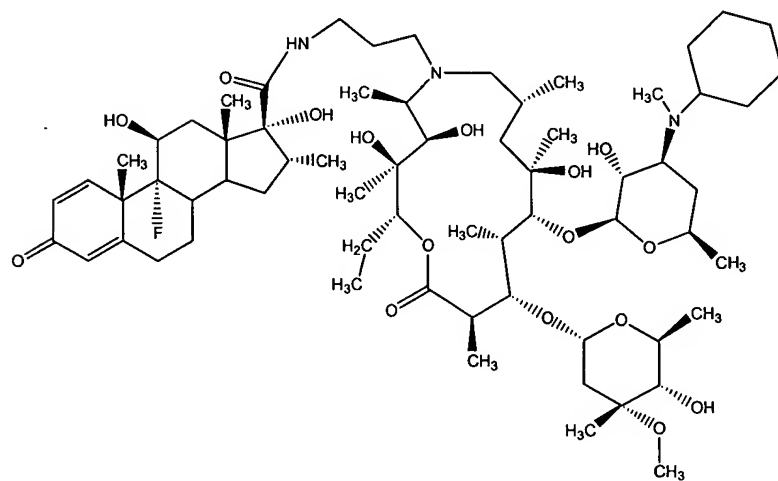
or a pharmaceutically acceptable salt or solvate thereof.

18. (Previously Presented) A compound of the formula



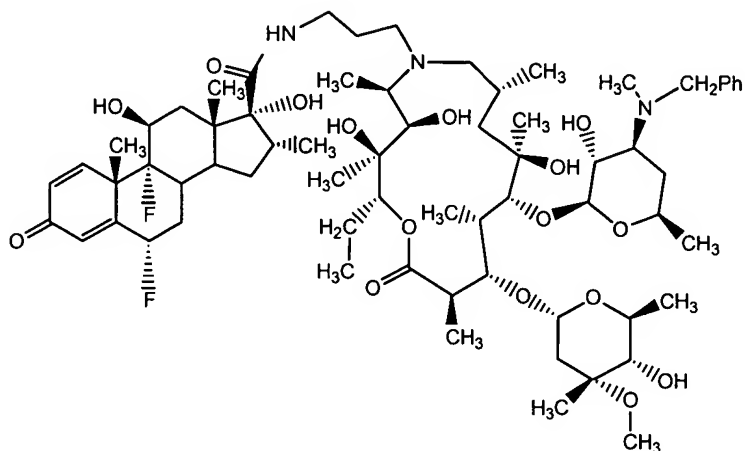
or a pharmaceutically acceptable salt or solvate thereof.

19. (Previously Presented) A compound of the formula



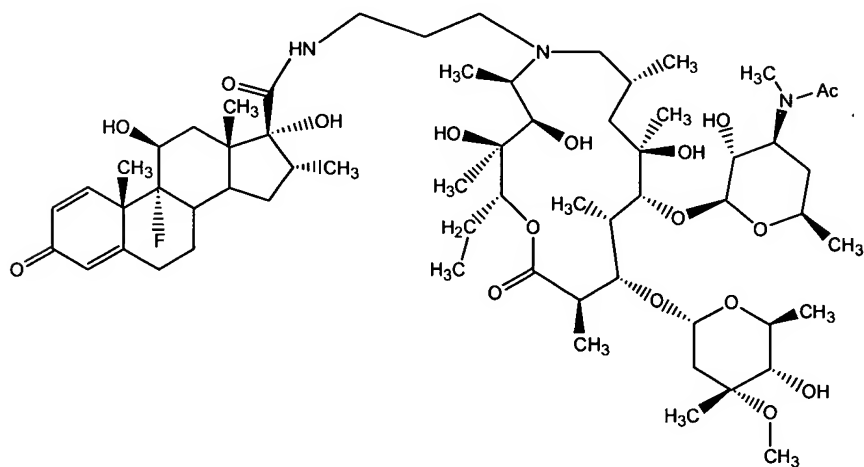
or a pharmaceutically acceptable salt or solvate thereof.

20. (Previously Presented) A compound of the formula



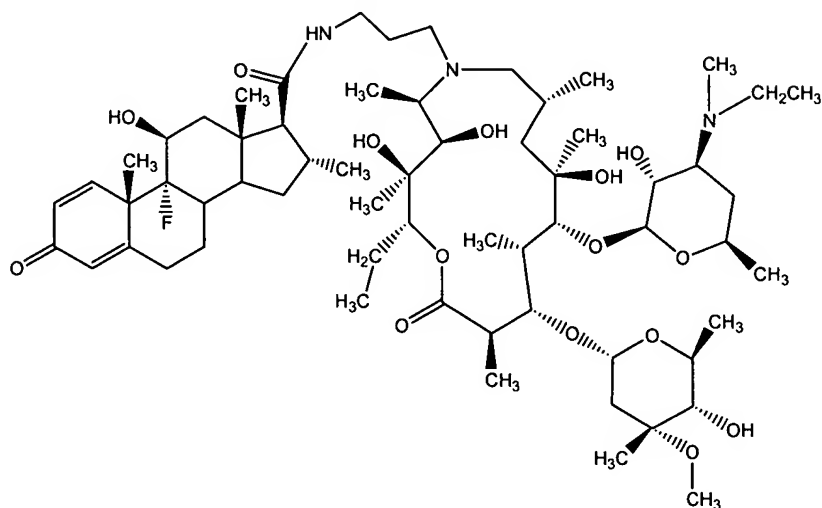
or a pharmaceutically acceptable salt or solvate thereof.

21. (Previously Presented) A compound of the formula



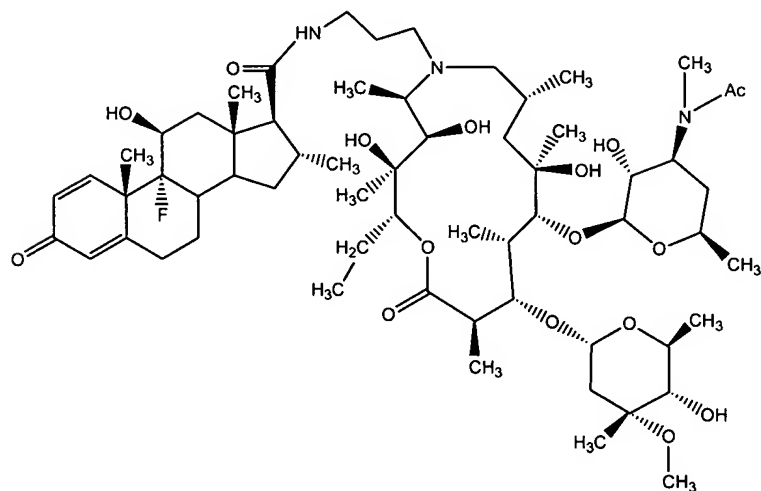
or a pharmaceutically acceptable salt or solvate thereof.

24. (Previously Presented) A compound of the formula



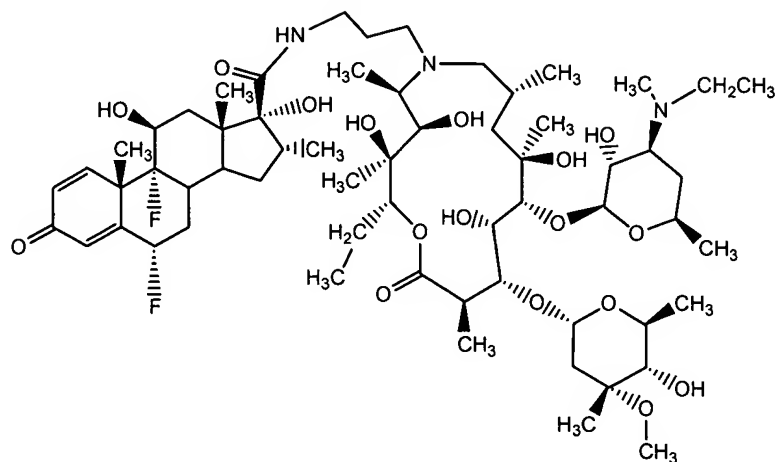
or a pharmaceutically acceptable salt or solvate thereof.

25. (Previously Presented) A compound of the formula



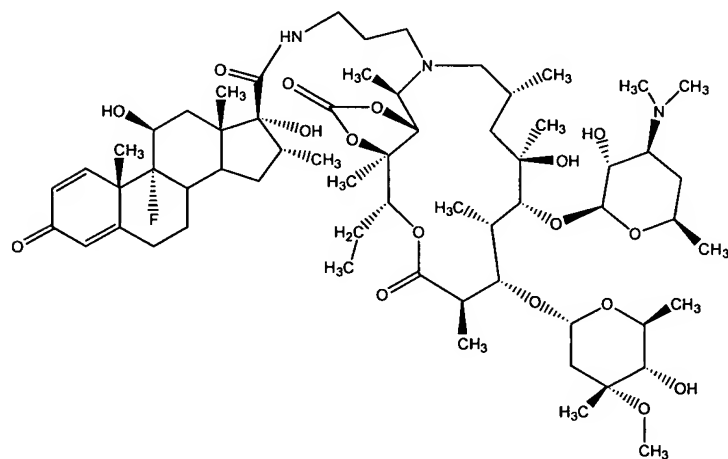
or a pharmaceutically acceptable salt or solvate thereof.

26. (Previously Presented) A compound of the formula



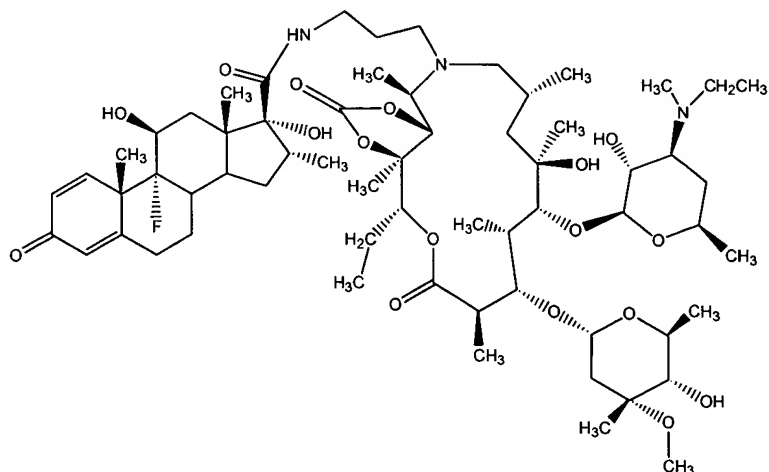
or a pharmaceutically acceptable salt or solvate thereof.

27. (Previously Presented) A compound of the formula



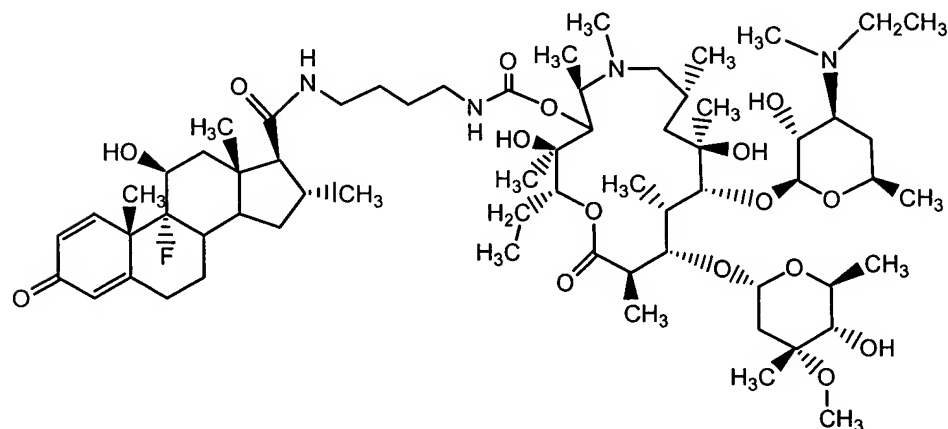
or a pharmaceutically acceptable salt or solvate thereof.

28. (Previously Presented) A compound of the formula



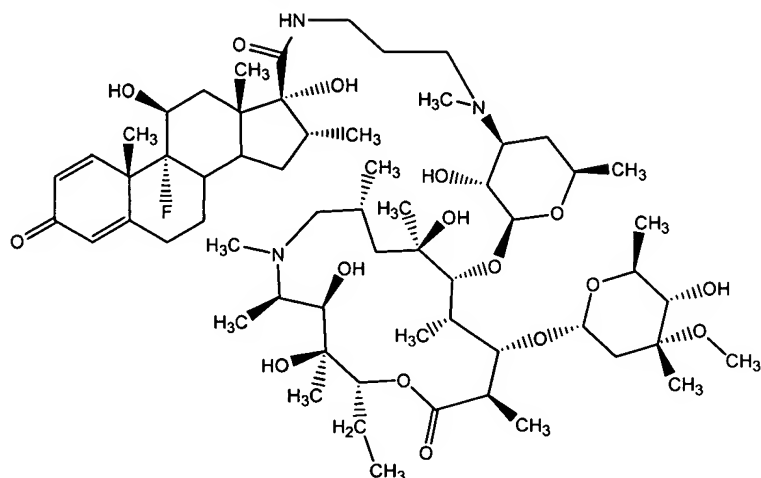
or a pharmaceutically acceptable salt or solvate thereof.

29. (Previously Presented) A compound of the formula



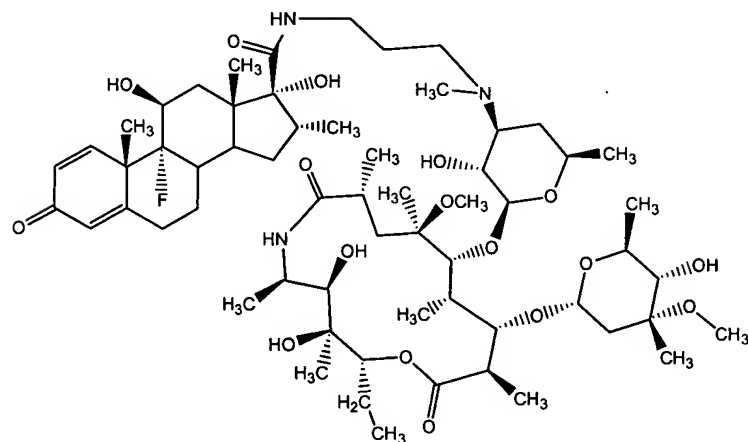
or a pharmaceutically acceptable salt or solvate thereof.

30. (Previously Presented) A compound of the formula



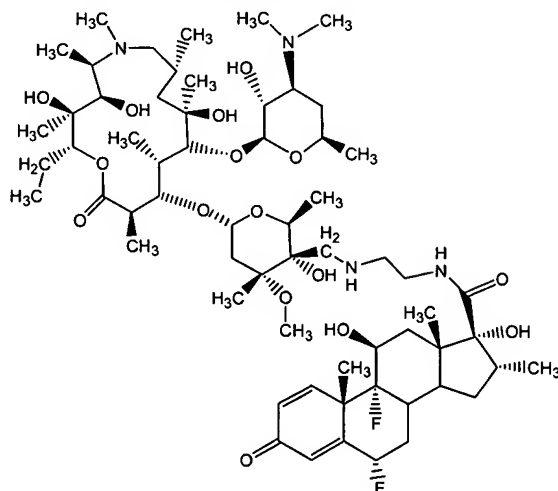
or a pharmaceutically acceptable salt or solvate thereof.

31. (Previously Presented) A compound of the formula



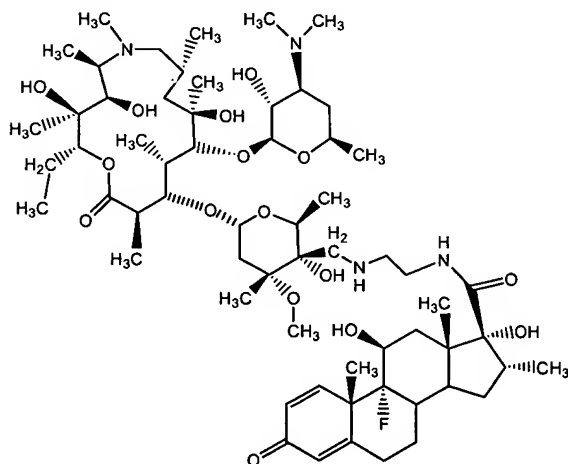
or a pharmaceutically acceptable salt or solvate thereof.

32. (Previously Presented) A compound of the formula



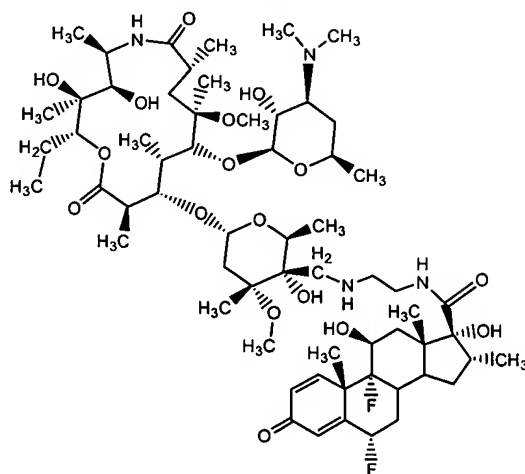
or a pharmaceutically acceptable salt or solvate thereof.

33. (Previously Presented) A compound of the formula



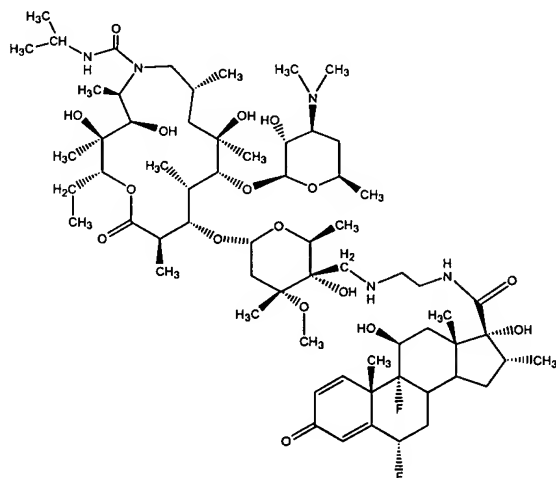
or a pharmaceutically acceptable salt or solvate thereof.

34. (Previously Presented) A compound of the formula



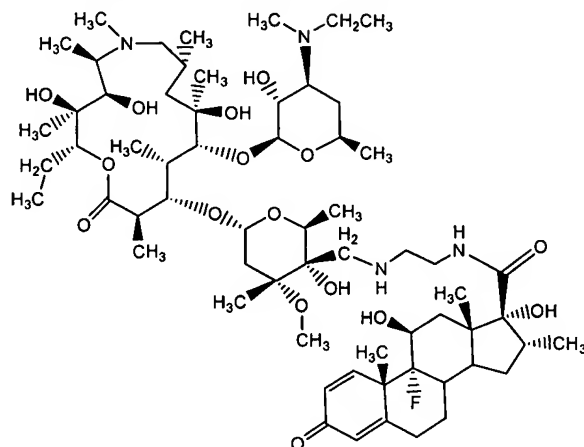
or a pharmaceutically acceptable salt or solvate thereof.

35. (Previously Presented) A compound of the formula



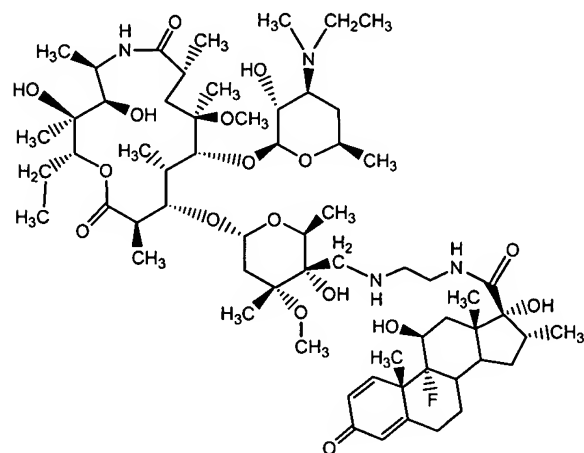
or a pharmaceutically acceptable salt or solvate thereof.

36. (Previously Presented) A compound of the formula



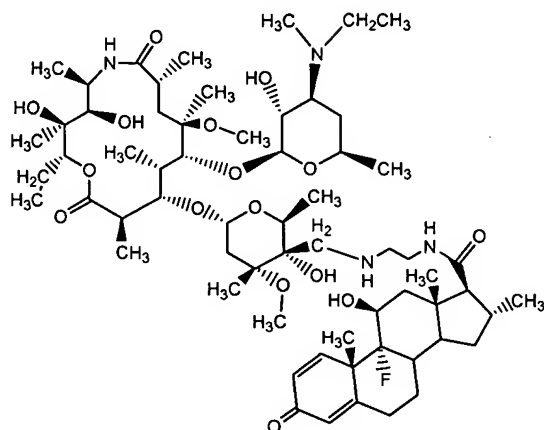
or a pharmaceutically acceptable salt or solvate thereof.

37. (Previously Presented) A compound of the formula



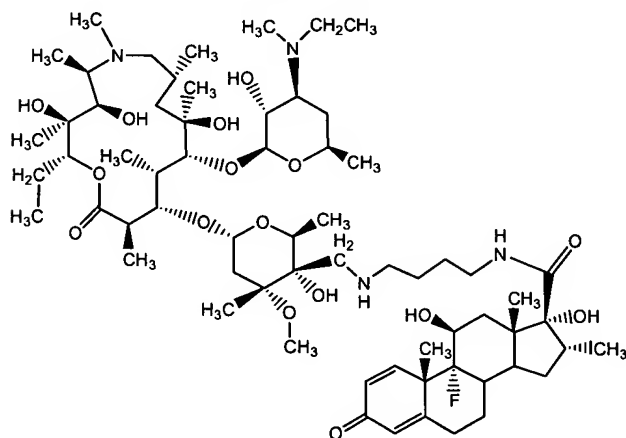
or a pharmaceutically acceptable salt or solvate thereof.

38. (Previously Presented) A compound of the formula



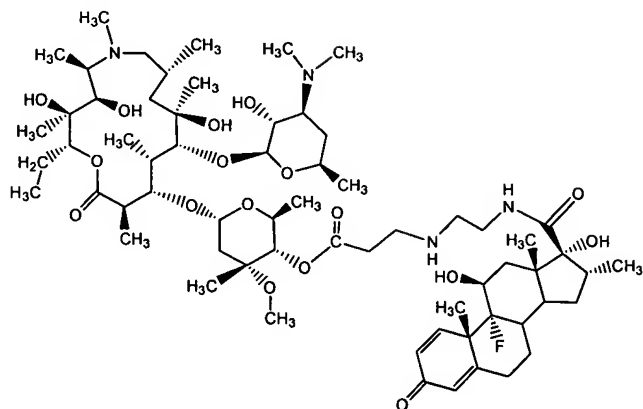
or a pharmaceutically acceptable salt or solvate thereof.

39. (Previously Presented) A compound of the formula



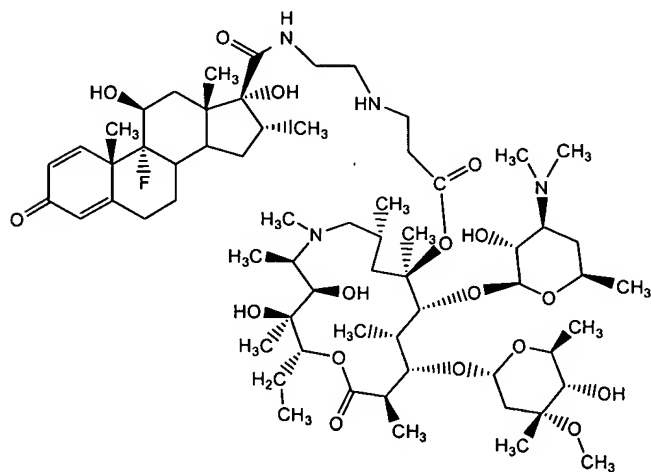
or a pharmaceutically acceptable salt or solvate thereof.

40. (Previously Presented) A compound of the formula



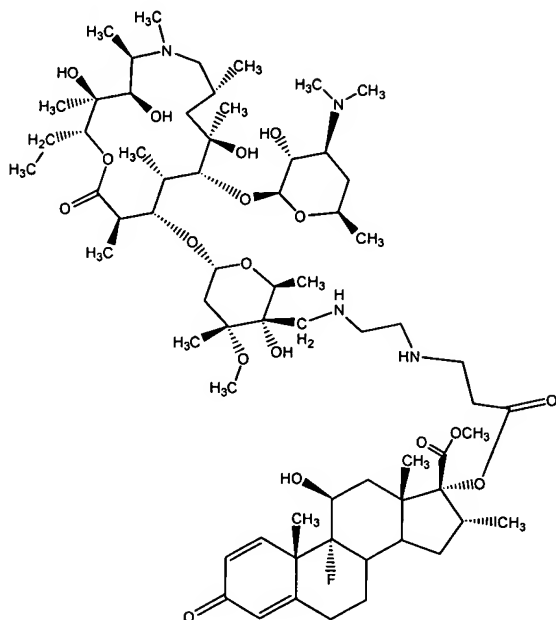
or a pharmaceutically acceptable salt or solvate thereof.

41. (Previously Presented) A compound of the formula



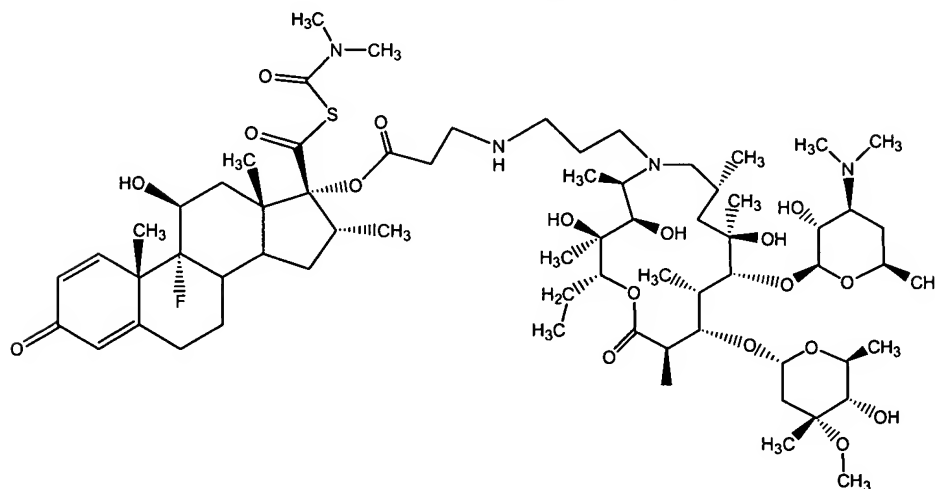
or a pharmaceutically acceptable salt or solvate thereof.

46. (Previously Presented) A compound of the formula



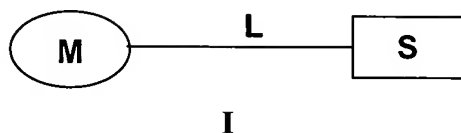
or a pharmaceutically acceptable salt or solvate thereof.

47. (Previously Presented) A compound of the formula

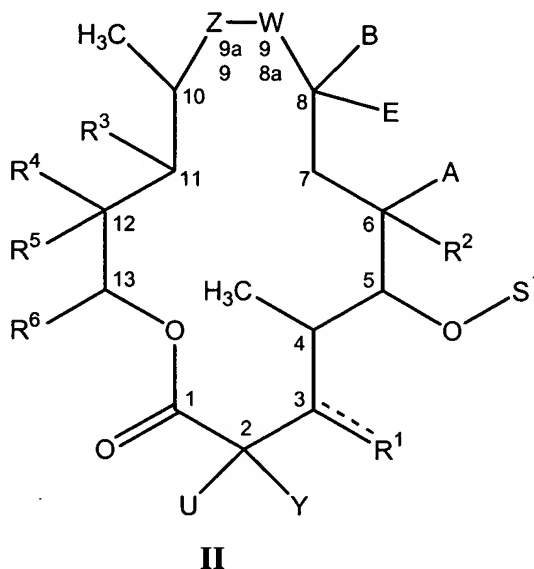


or a pharmaceutically acceptable salt or solvate thereof.

48. (Currently Amended) A process for the preparation for a compound of Formula I:



wherein **M** represents a group of Formula II:



wherein

- (i) Z and W independently are $>C=O$, $>CH_2$, $>CH-NR_tR_s$, $>N-R_N$ or $>C=N-R_M$, wherein

R_t and R_s independently are hydrogen or alkyl;

R_M is hydroxy, alkoxy, substituted alkoxy or OR^P ;

R_N is hydrogen, R^P, alkyl, alkenyl, alkynyl, alkoxy, alkoxyalkyl, or -C(X)-NR_tR_s; wherein X is =O or =S;

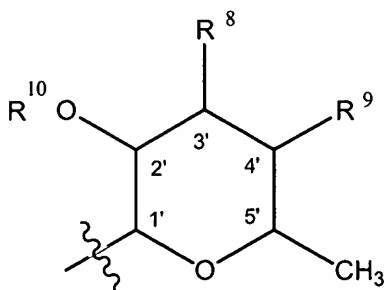
provided that Z and W cannot both simultaneously be, $>\text{C}=\text{O}$, $>\text{CH}_2$,

$$>\text{CH-NR}_t\text{R}_s, >\text{N-R}_N, >\text{C=N-R}_M \text{ or a bond};$$

- (ii) U and Y independently are hydrogen, halogen, alkyl, or hydroxyalkyl;

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(iii) R¹ is hydroxy, OR^p, -O-S² group or an =O;
(iv) S¹ is a sugar moiety of Formula **III**:



III

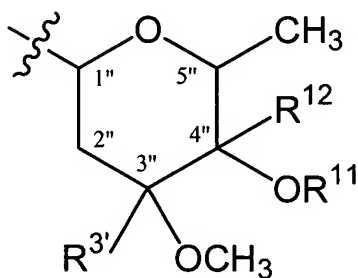
wherein

R⁸ and R⁹ are both hydrogen or together form a bond, or R⁹ is hydrogen and R⁸ is -N(CH₃)R^y, wherein

R^y is R^p, R^z or -C(O)R^z, wherein R^z is hydrogen or alkyl or alkenyl or alkynyl or cycloalkyl or aryl or heteroaryl or alkyl substituted with C₂-C₇-alkyl, C₂-C₇-alkenyl, C₂-C₇-alkynyl, aryl or heteroaryl;

R^{10} is hydrogen or R^p ;

S² sugar moiety of Formula **IV**:



IV

wherein

R^{3'} is hydrogen or methyl;

R¹¹ is hydrogen, R^p, or O-R¹¹ is a group that with R¹² and with C/4" carbon atom forms a >C=O or epoxy group;

R¹² is hydrogen or a group that with O-R¹¹ group and with C/4" carbon atom forms a >C=O or epoxy group;

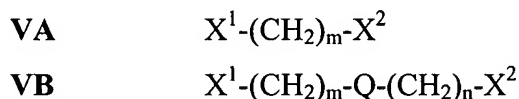
- (vi) R² is hydrogen, hydroxy, OR^P or alkoxy ;
- (vii) A is hydrogen or methyl;
- (viii) B is methyl or epoxy;
- (ix) E is hydrogen or halogen;
- R³ is hydroxy, OR^P , alkoxy or R³ is a group that with R⁵ and with C/11 and C/12 carbon atoms forms a cyclic carbonate or carbamate, or if W or Z is >N-R_N R³ is a group that with W or Z forms a cyclic carbamate;
- (xi) R⁴ is C₁-C₄ alkyl;
- (xii) R⁵ is hydrogen, hydroxy, OR^P , C₁-C₄ alkoxy, or a group that with R³ and with C/11 and C/12 carbon atoms forms a cyclic carbonate or carbamate;
- (xiii) R⁶ is hydrogen or C₁-C₄-alkyl; and
- R^P is hydroxyl or amino protective group;

wherein **M** has a linkage site through which it is linked to **S** *via* linking group **L**; provided that the linkage site being is at one or more of the following:

- any reactive hydroxy, nitrogen, or epoxy group located on S^1 , S^2 , or an aglycone oxygen if S^1 and/or S^2 is cleaved off;
- a reactive $>N-R_N$ or $-NR_tR_s$ or oxo group located on Z or W;
- a reactive hydroxy group located at any one of R^1 , R^2 , R^3 , and R^5 ;
- any other group that can be first derivatized to a hydroxy or $-NR_tR_s$ group; and

~~R^p is hydroxyl or amino protective group.~~

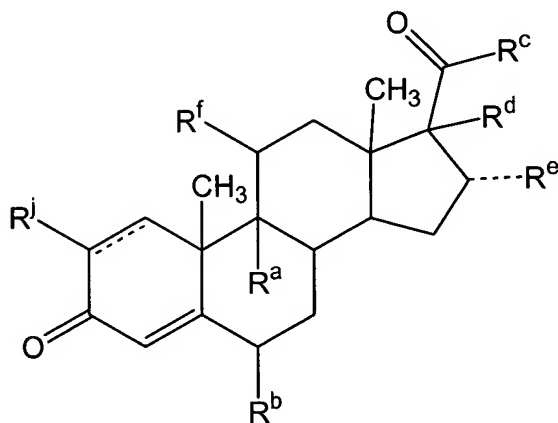
L represents a group of Formula **VA** or of Formula **VB**:



wherein

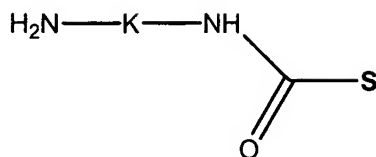
X¹ is selected from: -CH₂-, -CH₂NH-, -C(O)-, -OC(O)-, =N-O- or -OC(O)NH-;
-C(O)NH;

S represents a group of Formula X:



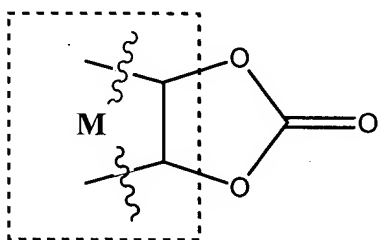
X

a) for a compound represented by Formula I comprising the steps of:



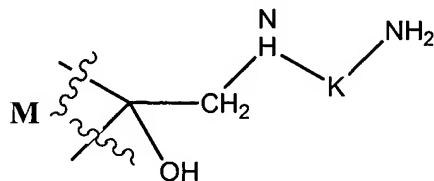
IVc

d) for a compound represented by Formula **I**, where X¹ is -OC(O)NH- and X² is -NHC(O)-, by reacting a macrolide represented by Formula **VII** and a free amino group of Formula **IVc**:



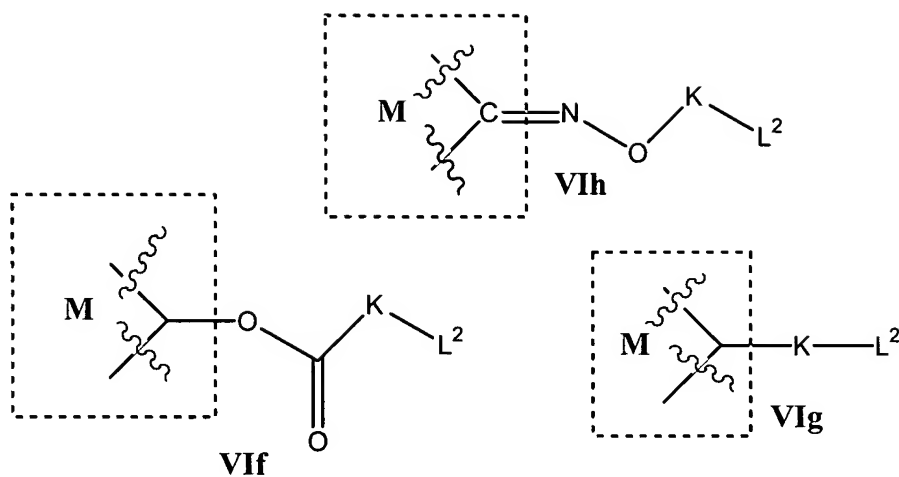
VII

e) for a compound represented by Formula I, X¹ is -CH₂-, Q is -NH- and X² is -NHC(O)-, by reacting a macrolide represented by Formula Va and a compound of Formula V:

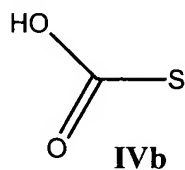


Va

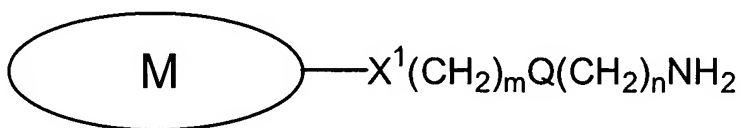
f) compound of Formula **I** by reacting a macrolide represented by Formula **VI**f or by Formula **VI**g or by Formula **VI**h having a leaving group L_2



with a free carboxyl acid of steroid represented by Formula IVb

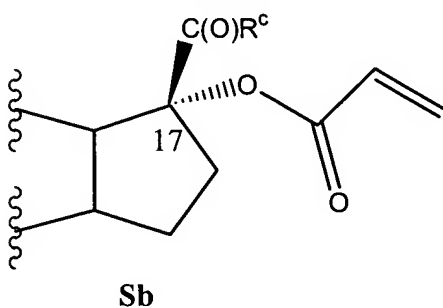


g) for a compound represented by Formula I, wherein X¹ is -OC(O)-, Q is NH and X² is -NH- by reacting a macrolide represented by:



VI d

and a steroid subunit having a -C=C- bond represented by Formula Sb:



following by modification of R^c group.

49. (Previously Presented) A pharmaceutical composition comprising a compound according to claim 1 or a pharmaceutically acceptable salt or solvate thereof as well as pharmaceutically acceptable diluent or carrier.
50. (Previously Presented) A method of treatment of inflammatory diseases, disorders or conditions characterized by or associated with an undesirable inflammatory immune response, and all diseases and conditions induced by or associated with an excessive secretion of TNF- α and IL-1 which comprises administering to a subject in need of treatment a therapeutically effective amount of a compound according to claim 1.
51. (Previously Presented) A method of treating inflammatory conditions or immune or anaphylactic disorders associated with infiltration of leukocytes into inflamed tissue in a subject in need thereof which comprises administering to said subject a therapeutically effective amount of a compound according to claim 1.
52. (Previously presented) The method according to claim 51, wherein inflammatory conditions and immune disorders are selected from the group consisting of asthma, adult respiratory distress syndrome, bronchitis, and cystic fibrosis.
53. (Previously presented) A method according to claim 51, wherein said inflammatory conditions and immune disorders are selected from the group consisting of inflammatory conditions or immune disorders of the lungs, joints, eyes, bowel, skin, and heart.
54. (Previously presented) A method according to claim 51, wherein said inflammatory conditions and immune disorders are selected from the group consisting of asthma, adult respiratory distress syndrome,

bronchitis, cystic fibrosis, rheumatoid arthritis, rheumatoid spondylitis, osteoarthritis, gouty arthritis, uveitis, conjunctivitis, inflammatory bowel conditions, Crohn's disease, ulcerative colitis, distal proctitis, psoriasis, eczema, dermatitis, coronary infarct damage, chronic inflammation, endotoxin shock, and smooth muscle proliferation disorders.

55. (Previously Presented) A pharmaceutical composition comprising a compound according to claim 2, or a pharmaceutically acceptable salt or solvate thereof as well as pharmaceutically acceptable diluent or carrier.
56. (Previously Presented) A method of treatment of inflammatory diseases, disorders or conditions characterized by or associated with an undesirable inflammatory immune response, and all diseases and conditions induced by or associated with an excessive secretion of TNF- α and IL-1 which comprises administering to a subject in need of treatment a therapeutically effective amount of a compound according to claim 2.
57. (Previously Presented) A method of treating inflammatory conditions or immune or anaphylactic disorders associated with infiltration of leukocytes into inflamed tissue in a subject in need thereof which comprises administering to said subject a therapeutically effective amount of a compound according to claim 2.
58. (Previously Presented) The method according to claim 57, wherein inflammatory conditions and immune disorders are selected from the group consisting of asthma, adult respiratory distress syndrome, bronchitis, and cystic fibrosis.
59. (Previously Presented) A method according to claim 57, wherein said inflammatory conditions and immune disorders are selected from the

group consisting of inflammatory conditions or immune disorders of the lungs, joints, eyes, bowel, skin, and heart.

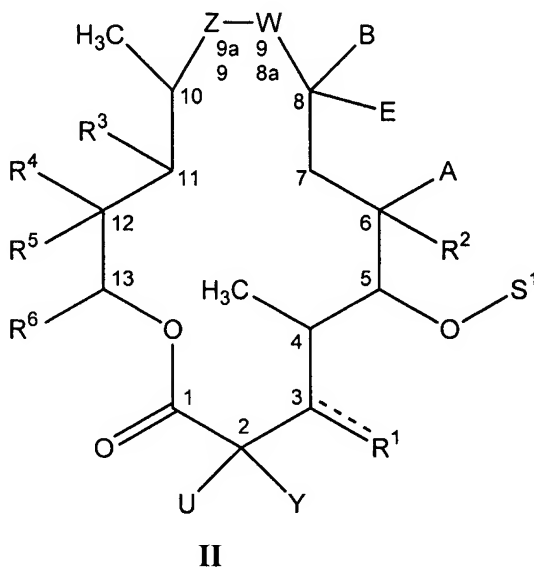
60. (Previously Presented) A method according to claim 57, wherein said inflammatory conditions and immune disorders are selected from the group consisting of asthma, adult respiratory distress syndrome, bronchitis, cystic fibrosis, rheumatoid arthritis, rheumatoid spondylitis, osteoarthritis, gouty arthritis, uveitis, conjunctivitis, inflammatory bowel conditions, Crohn's disease, ulcerative colitis, distal proctitis, psoriasis, eczema, dermatitis, coronary infarct damage, chronic inflammation, endotoxin shock, and smooth muscle proliferation disorders.

61. (New) A compound of the formula:



I

wherein **M** represents a group of Formula **II**:

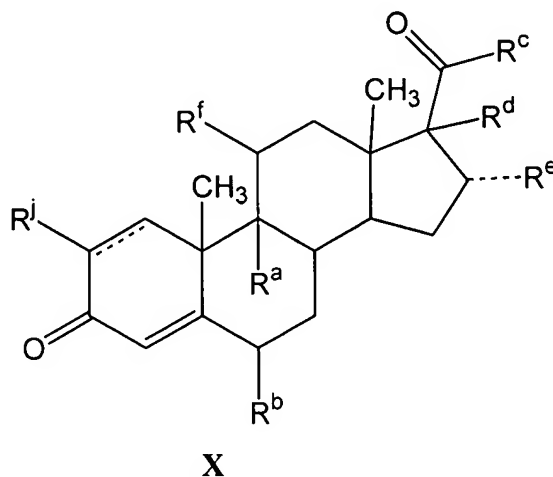


c) a reactive hydroxy group located at any one of R^1 , R^2 , R^3 , and R^5 ;

d) any other group that can be first derivatized to a hydroxy or

$-NR_iR_s$ group; and

S represents a group of Formula X:



wherein

R^a and R^b independently represents, hydrogen or halogen;

R^c is a valence-bond;

R^d and R^e independently represents: hydrogen, hydroxy, methyl or C_1 - C_4 -alkoxy or each are a group that forms a 1,3-dioxolane ring with the other;

R^f is hydrogen, hydroxy, chloro, or forming a keto group with the carbon atom it is attached to;

R^j is hydrogen or chloro;

or a pharmaceutically acceptable salt or solvate thereof;

wherein

L is a linker molecule to which each of M and S are covalently linked.

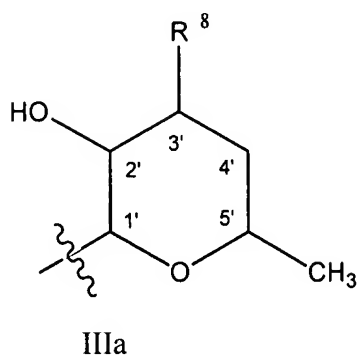
62. (New) The compound according to claim 61 wherein

(i) Z is $>N-R_N$ and W is $>CH_2$;

(ii) U is H and Y is CH_3 ;

(iii) R^1 is an $-O-S^2$ group;

(iv) S^1 is a sugar moiety of Formula **IIIa**:

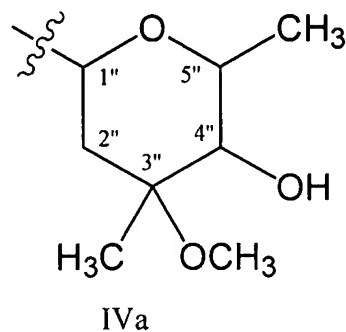


wherein

R^8 is H or $-N(CH_3)R^y$, wherein

R^y an alkyl;

S^2 sugar moiety of Formula **IVa**:



(vi) R^2 is hydroxy;

(vii) A is CH_3 ;

(viii) B is CH_3 ;

(ix) E is H;

(x) R^3 is OH;

(xi) R^4 is C_1 - C_4 alkyl;

(xii) R^5 is OH; and

(xiii) R^6 is C_1 - C_4 -alkyl;

R^d is H or OH;

or a pharmaceutically acceptable salt or solvate thereof.